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 FILE 'REGISTRY' ENTERED AT 16:25:56 ON 10 JUL 2001  
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STRUCTURE FILE UPDATES: 9 JUL 2001 HIGHEST FN 345196-14-7  
 DICTIONARY FILE UPDATES: 9 JUL 2001 HIGHEST FN 345196-14-7

TICA INFORMATION NOW CURRENT THROUGH January 11, 2001

Please note that search-term pricing does apply when  
 conducting SmartSELECT searches.

Structure search limits have been increased. See HELP SLIMIT  
 for details.

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 1.4 STR  

$$\begin{array}{ccccc} 1 & C & N & 3 & \\ & & & & 10 \\ & & & & 6 \\ 6 & C & 4 & C & \\ 1 & \text{G1} & 3 & N & C & N \\ & 5 & 7 & 8 & 9 & \end{array}$$

Point of Contact:  
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 CM11201 Tel: 308-4498

VAR G1=X/NO2/N/C/CY

NODE ATTRIBUTES:

NSPEC IS EC AT +  
 DEFAULT MLEVEL IS ATOM  
 DEFAULT ELEVEL IS LIMITED

GRAPH ATTRIBUTES:

FIG(S) ARE ISOLATED OR EMBEDDED  
 NUMBER OF NODES IS 10

STEREO ATTRIBUTES: NONE

L1 29632 SEA FILE=REGISTRY ABB=ON PLU=ON NCSC?/ES  
 L7 1667 SEA FILE=REGISTRY SUB=L5 SSS FUL L4  
 L8 1250 SEA FILE=REGISTRY ABB=ON PLU=ON L7 AND (16.299.11 OR  
 16.299.12)/RID

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(FILE 'REGISTRY' ENTERED AT 16:11:49 ON 10 JUL 2001)  
 SAV GERSTL&07 A L7

L8 1250 S L7 AND (16.299.11 OR 16.299.12)/RID  
 L9 417 S L7 NOT L8

FILE 'HCAPLUS' ENTERED AT 16:13:43 ON 10 JUL 2001

L10 134 S L8  
 L11 26 S L8 (L) THU/FL  
 L12 47 S L8 (L) BAC/FL  
 L13 57 S L8 AND (1 OR 63)/SC, SX  
 L14 74 S L11-L13  
 L15 106 S L10 AND (PY=1998 OR PY<=1995 OR AY=1998)  
 L16 61 S L15 AND L14  
 L17 36 S L15 AND (?INFOFLAS? OR ?TUMOR? OR ?TUMOUR? OR ?CANCER? OR ?CAR  
 L18 4 S L15 AND (?PREGNIFERAT? OR ?CYTOTOX?)  
 L19 58 S L17, L18  
 L20 22 S L19 NOT (1 OR 63)/SC, SX  
 L21 3 S L20 NOT 4/SC  
 L22 2 S L21 NOT 1/SC  
 L23 15 S L19 NOT L20

L24 11 S L23 NOT 4/SC  
 L25 6 S L14 AND (FAF OF LEWIS OR CYCLIN OR INOSINE OR ANTITUMOR) /TI  
 L26 8 S L12, L25  
 L27 47 S L16 NOT L17  
 L28 45 S L7 NOT 4/SC

FILE 'REGISTRY' ENTERED AT 16:25:56 ON 10 JUL 2001

=. fil hcaplus

FILE 'HCAPLUS' ENTERED AT 16:26:10 ON 10 JUL 2001

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FILE COVERS 1947 - 10 Jul 2001 VOL 135 ISS 3

FILE LAST UPDATED: 9 Jul 2001 (20010709/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

This file supports REGISTRY for direct browsing and searching of all substance data from the REGISTRY file. Enter HELP FIRST for more information.

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=. d 126 bib abs hitrn fhitstr tot

LN:6 ANSWER 1 OF 8 HCAPLUS COPYRIGHT 2001 ACS  
 AN 2001:111513 HCAPLUS  
 DE 134:163040  
 TI Preparation of heteroaryl aryl ureas as raf kinase inhibitors  
 IN Wood, Jili E.; Wild, Hanne; Rogers, Daniel H.; Lyons, John; Katz, Michael;  
 Caringal, Yolanda; Dally, Robert; Lee, Wendy; Smith, Roger A.; Blum, Cheri  
 PA Onyx Pharmaceuticals, USA; Bayer Corporation  
 SC U.S., 30 pp.  
 CODEN: USXXAM

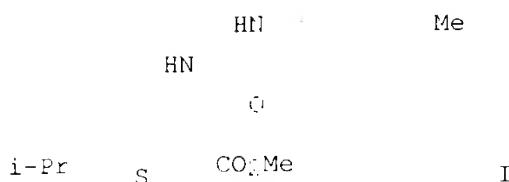
DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 6187799	B1	20010213	US 1948-83399	19980522 <--
	US 2001006975	A1	20010705	US 2001-755060	20010108 <--
PKAI	US 1997-126420	P	19970523 <--		
	US 1998-83399	A3	19980522 <--		

GI



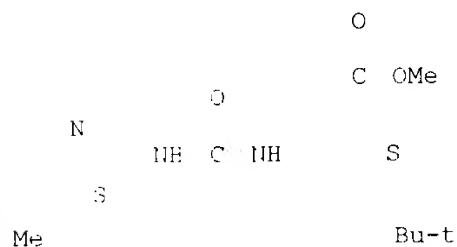
AB The title heteroaryl aryl ureas, useful in treating **tumors** mediated by raf kinase (no data), were prep'd. E.g., a multi-step synthesis of the urea I was given. The title compds. such as I are effective at 0.01-200 mg/kg/day.

IT **216589-90-1P**  
 PL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
 (prepn. of heteroaryl aryl ureas as raf kinase inhibitors)

IT **216589-90-1P**  
 PL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
 (prepn. of heteroaryl aryl ureas as raf kinase inhibitors)

RN 216589-90-1 HCPLUS

CN 2-Thiophenecarboxylic acid, 5-(1,1-dimethylethyl)-3-[[[(5-methyl-2-thiazolyl)amino]carbonyl]amino]-, methyl ester (9CI) (CA INDEX NAME)



RE.CNT 36

RE  
 (1) Acker; US 4437878 1984 HCPLUS  
 (2) Aldrich; US 4009847 1977 HCPLUS  
 (3) Anon; JP 54-32468 1979 HCPLUS  
 (4) Anon; DE 3305866 1984 HCPLUS  
 (5) Anon; WO 9324458 1993 HCPLUS  
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

LC6 ANSWER 2 OF 8 HCPLUS COPYRIGHT 2001 ACS

AN 2000:314688 HCPLUS

DN 132:334455

TI 2-Ureidothiazole derivatives, process for their preparation, and their use as **antitumor** agents

IN Pevarello, Paolo; Amici, Raffaella; Traquandi, Gabriella; Villa, Manuela; Vulpetti, Anna; Isacchi, Antonella

PA Pharmacia & Upjohn S.p.A., Italy

SO PCT Int. Appl., 95 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

PATENT NO.

KIND DATE

APPLICATION NO. DATE

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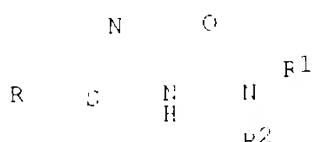
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PI WO 2000026203 A1 20000511 WO 1994-EP8307 19991027 <--  
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 IL, IN, IS, JP, KP, KE, LC, LF, LR, LT, LV, MG, MK, MN, MX, NO,  
 NZ, PL, RO, SG, SI, SF, SL, TF, TT, UA, US, UZ, VN, YU, ZA, AM,  
 AZ, BY, EG, KZ, MD, RU, TJ, TM  
 FW: GH, GM, GE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,  
 DK, ES, FI, FR, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,  
 CG, CI, CM, GA, GN, GW, ML, MF, NE, SN, TI, TG

PRAI GB 1998-00873 A 19981030 &lt;--

OS CARPAT 130:334455

GI



AB The title 1-ureido-1,3-thiazole derivs. I and their pharmaceutically acceptable salts are disclosed [wherein R = halo, nitro, (un)substituted amino, C1-6 alkyl, C3-6 cycloalkyl, aryl, or arylalkyl; P1 = (un)substituted C1-6 alkyl, 3- to 6-membered carbocycle or 5- to 7-membered heterocycle, aryl, arylcarbonyl, or arylalkyl; P2 = H, straight or branched C1-4 alkyl, C2-4 alkenyl, or alkynyl; or NF1F2 = (un)substituted, optionally benco-condensed or bridged 5- to 7-membered heterocycle, or 9- to 11-membered spiro-heterocycle]. The compds. are active as cdk/cyclin inhibitors, and are useful for treating cell proliferative disorders assocd. with an altered cell dependent kinase activity. The **proliferative** disorders include **cancer** and a wide variety of other conditions, such as Alzheimer's disease, viral infections, autoimmune diseases, and neurodegenerative disorders. Over 230 invention compds. are claimed and/or prepd. in examples. For instance, reaction of Pt. isocyanate with 2-amino-5-nitro-1,3-thiazole hydrotrionide in the presence of Et3N gave title compd. I [R = Pr, P1 = Ph, R' = H]. The similarly prepd. title compd. I [R = iso-Pr, P1 = 3,5-dimethylphenyl, R' = H] inhibited cdk2/cyclin A complex in vitro with an IC50 of 0.56 .mu.M.

IT 267431-26-5, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2-nitrophenyl)urea  
 FL: RCT (Reactant)  
 (starting material; prepn. of ureidothiazole derivs. as **antitumor** agents)  
 IT 267429-35-6P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-nitrophenyl)urea 267429-43-6P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3-nitrophenyl)urea 267429-47-0P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3-aminophenyl)urea 267431-00-5P, N-(3-Isopropyl-1,3-thiazol-2-yl)-N'-(3-aminophenyl)urea  
 FL: BAC (Biological activity or effector, except adverse); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PEP (Preparation); USES (Uses);  
 (target compd.; prepn. of ureidothiazole derivs. as **antitumor** agents)

IT 14954-34-8P, N-(5-Methyl-1,3-thiazol-2-yl)-N'-phenylurea  
 202056-91-5P, N-(5-Phenyl-1,3-thiazol-2-yl)-N'-(4-chlorophenyl)urea 267428-92-2P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-phenylurea 267428-93-3P, N-(5-Bromo-1,3-thiazol-2-yl)-N'-phenylurea 267428-94-4P, N-(5-Phenyl-1,3-thiazol-2-yl)-N'-phenylurea 267428-95-5P, N-(5-Cyclopropyl-1,3-thiazol-2-yl)-N'-phenylurea 267428-96-6P, N-(5-Bromo-1,3-thiazol-2-yl)-N'-(4-sulfamoylphenyl)urea 267428-97-7P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-sulfamoylphenyl)urea 267428-98-8P, N-(5-Phenyl-1,3-thiazol-2-yl)-N'-(4-sulfamoylphenyl)urea 267428-99-9P, N-(5-Cyclopropyl-1,3-thiazol-2-yl)-N'-(4-

sulfamoylphenyl)urea **267429-00-5P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3-methoxyphenyl)urea **267429-01-6P**, N-(5-Bromo-1,3-thiazol-2-yl)-N'-(3-methoxyphenyl)urea **267429-02-7P**, N-(5-Phenyl-1,3-thiazol-2-yl)-N'-(3-methoxyphenyl)urea **267429-03-8P**, N-(5-Cyclopropyl-1,3-thiazol-2-yl)-N'-(3-methoxyphenyl)urea **267429-04-9P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-chlorophenyl)urea **267429-05-0P**, N-(5-Bromo-1,3-thiazol-2-yl)-N'-(4-chlorophenyl)urea **267429-06-1P**, N-(5-Cyclopropyl-1,3-thiazol-2-yl)-N'-(4-chlorophenyl)urea **267429-07-2P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3-chlorophenyl)urea **267429-08-3P**, N-(5-Bromo-1,3-thiazol-2-yl)-N'-(3-chlorophenyl)urea **267429-09-4P**, N-(5-Phenyl-1,3-thiazol-2-yl)-N'-(3-chlorophenyl)urea **267429-10-7P**, N-(5-Cyclopropyl-1,3-thiazol-2-yl)-N'-(3-chlorophenyl)urea **267429-11-8P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2-chlorophenyl)urea **267429-12-9P**, N-(5-Bromo-1,3-thiazol-2-yl)-N'-(2-chlorophenyl)urea **267429-13-0P**, N-(5-Phenyl-1,3-thiazol-2-yl)-N'-(2-chlorophenyl)urea **267429-14-1P**, N-(5-Cyclopropyl-1,3-thiazol-2-yl)-N'-(2-chlorophenyl)urea **267429-15-2P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-methoxyphenyl)urea **267429-16-3P**, N-(5-Bromo-1,3-thiazol-2-yl)-N'-(4-methoxyphenyl)urea **267429-17-4P**, N-(5-Phenyl-1,3-thiazol-2-yl)-N'-(4-methoxyphenyl)urea **267429-18-5P**, N-(5-Cyclopropyl-1,3-thiazol-2-yl)-N'-(4-methoxyphenyl)urea **267429-19-6P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-hydroxyphenyl)urea **267429-20-9P**, N-(5-Bromo-1,3-thiazol-2-yl)-N'-(4-hydroxyphenyl)urea **267429-21-0P**, N-(5-Phenyl-1,3-thiazol-2-yl)-N'-(4-hydroxyphenyl)urea **267429-22-1P**, N-(5-Cyclopropyl-1,3-thiazol-2-yl)-N'-(4-hydroxyphenyl)urea **267429-23-2P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3-hydroxyphenyl)urea **267429-24-3P**, N-(5-Bromo-1,3-thiazol-2-yl)-N'-(3-hydroxyphenyl)urea **267429-25-4P**, N-(5-Phenyl-1,3-thiazol-2-yl)-N'-(3-hydroxyphenyl)urea **267429-26-5P**, N-(5-Cyclopropyl-1,3-thiazol-2-yl)-N'-(3-hydroxyphenyl)urea **267429-27-6P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2-methoxyphenyl)urea **267429-28-7P**, N-(5-Bromo-1,3-thiazol-2-yl)-N'-(2-methoxyphenyl)urea **267429-29-8P**, N-(5-Phenyl-1,3-thiazol-2-yl)-N'-(2-methoxyphenyl)urea **267429-30-1P**, N-(5-Cyclopropyl-1,3-thiazol-2-yl)-N'-(2-methoxyphenyl)urea **267429-31-2P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2-hydroxyphenyl)urea **267429-32-3P**, N-(5-Bromo-1,3-thiazol-2-yl)-N'-(2-hydroxyphenyl)urea **267429-33-4P**, N-(5-Phenyl-1,3-thiazol-2-yl)-N'-(2-hydroxyphenyl)urea **267429-34-5P**, N-(5-Cyclopropyl-1,3-thiazol-2-yl)-N'-(2-hydroxyphenyl)urea **267429-36-7P**, N-(5-Bromo-1,3-thiazol-2-yl)-N'-(4-nitrophenyl)urea **267429-37-8P**, N-(5-Phenyl-1,3-thiazol-2-yl)-N'-(4-nitrophenyl)urea **267429-38-9P**, N-(5-Cyclopropyl-1,3-thiazol-2-yl)-N'-(4-nitrophenyl)urea **267429-39-0P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-aminophenyl)urea **267429-40-3P**, N-(5-Bromo-1,3-thiazol-2-yl)-N'-(4-aminophenyl)urea **267429-41-4P**, N-(5-Phenyl-1,3-thiazol-2-yl)-N'-(4-aminophenyl)urea **267429-42-5P**, N-(5-Cyclopropyl-1,3-thiazol-2-yl)-N'-(4-aminophenyl)urea **267429-44-7P**, N-(5-Bromo-1,3-thiazol-2-yl)-N'-(3-nitrophenyl)urea **267429-45-8P**, N-(5-Phenyl-1,3-thiazol-2-yl)-N'-(3-nitrophenyl)urea **267429-46-9P**, N-(5-Cyclopropyl-1,3-thiazol-2-yl)-N'-(3-nitrophenyl)urea **267429-48-1P**, N-(5-Bromo-1,3-thiazol-2-yl)-N'-(3-aminophenyl)urea **267429-49-2P**, N-(5-Phenyl-1,3-thiazol-2-yl)-N'-(3-aminophenyl)urea **267429-50-5P**, N-(5-Cyclopropyl-1,3-thiazol-2-yl)-N'-(3-aminophenyl)urea **267429-51-6P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-benzylurea **267429-52-7P**, N-(5-Bromo-1,3-thiazol-2-yl)-N'-benzylurea **267429-53-8P**, N-(5-Phenyl-1,3-thiazol-2-yl)-N'-benzylurea **267429-54-9P**, N-(5-Cyclopropyl-1,3-thiazol-2-yl)-N'-benzylurea **267429-55-0P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(pyrid-3-yl)urea **267429-56-1P**, N-(5-Bromo-1,3-thiazol-2-yl)-N'-(pyrid-3-yl)urea **267429-57-2P**, N-(5-Phenyl-1,3-thiazol-2-yl)-N'-(pyrid-3-yl)urea **267429-58-3P**, N-(5-Cyclopropyl-1,3-thiazol-2-yl)-N'-(pyrid-3-yl)urea

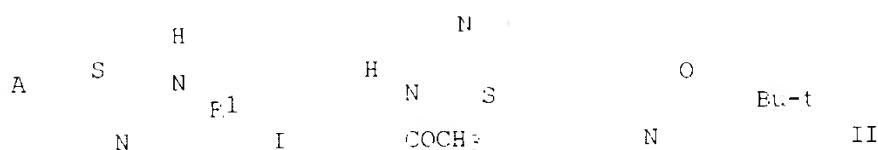
267429-59-4P, N-(5-Bromo-1,3-thiazol-2-yl)-N'-(pyrid-4-yl)urea  
 267429-60-7P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(pyrid-4-yl)urea  
 267429-61-8P, N-(5-Phenyl-1,3-thiazol-2-yl)-N'-(pyrid-4-yl)urea  
 267429-62-9P, N-(5-Cyclopropyl-1,3-thiazol-2-yl)-N'-(pyrid-4-yl)urea  
 267429-63-0P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(pyrid-2-yl)urea  
 267429-64-1P, N-(5-Bromo-1,3-thiazol-2-yl)-N'-(pyrid-2-yl)urea  
 267429-65-2P, N-(5-Phenyl-1,3-thiazol-2-yl)-N'-(pyrid-2-yl)urea  
 267429-66-3P, N-(5-Cyclopropyl-1,3-thiazol-2-yl)-N'-(pyrid-2-yl)urea  
 267429-67-4P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(benzothiophen-2-yl)urea  
 267429-68-5P, N-(5-Bromo-1,3-thiazol-2-yl)-N'-(benzothiophen-2-yl)urea  
 267429-69-6P, N-(5-Phenyl-1,3-thiazol-2-yl)-N'-(benzothiophen-2-yl)urea  
 267429-70-9P, N-(5-Isopropyl-1,3-thiazol-2-yl)-4-morpholinecarboxamide  
 267429-71-0P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-methylphenyl)urea  
 267429-72-1P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3-fluorophenyl)urea  
 267429-73-2P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-cyanophenyl)urea  
 267429-74-3P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3-cyanophenyl)urea  
 267429-75-4P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4,6-dimethylphenyl)urea  
 267429-76-5P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-fluorobenzyl)urea  
 267429-77-6P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3-acetylphenyl)urea  
 267429-78-7P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-acetylphenyl)urea  
 267429-79-8P, 3-[(5-Isopropyl-1,3-thiazol-2-yl)amino]benzoic acid  
 267429-80-1P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-isopropylphenyl)urea  
 267429-81-2P, 3-[(5-Isopropyl-1,3-thiazol-2-yl)amino]benzamide  
 267429-82-3P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-methoxybenzyl)urea  
 267429-83-4P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-butylphenyl)urea  
 267429-84-5P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-(trifluoromethyl)phenyl)urea  
 267429-85-6P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3-bromophenyl)urea  
 267429-86-7P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-cyclohexylphenyl)urea  
 267429-87-8P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-phenoxypyhenyl)urea  
 267429-88-9P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-benzoxypyhenyl)urea  
 267429-89-0P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4,5-dimethylphenyl)urea  
 267429-90-3P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4,4-dimethylphenyl)urea  
 267429-91-4P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3-methoxy-[1,1'-biphenyl]-4-yl)urea  
 267429-92-5P, N-(5-Isopropyl-1,3-thiazol-2-yl)-3,4-dihydro-2(1H)-isoquinolinecarboxamide  
 267429-93-6P, N-Benzyl-N'-(5-isopropyl-1,3-thiazol-2-yl)-N-methyliurea  
 267429-94-7P, N-(5-Isopropyl-1,3-thiazol-2-yl)-6,7-dimethoxy-3,4-dihydro-2(1H)-isoquinolinecarboxamide  
 267429-95-8P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3-chloro-4-methylphenyl)urea  
 267429-96-9P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3-chloro-6-methylphenyl)urea  
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 267429-99-2P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2-methoxy-5-chlorophenyl)urea  
 267430-00-2P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2-chloro-4-methoxyphenyl)urea  
 267430-01-3P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3,5-dichlorophenyl)urea  
 267430-02-4P, N-([1,1'-Biphenyl]-2-yl)-N'-(5-isopropyl-1,3-thiazol-2-yl)urea  
 267430-03-5P, N-Ethyl-N'-(5-isopropyl-1,3-thiazol-2-yl)-N-phenylurea  
 267430-04-6P, N-[4-[(5-Isopropyl-1,3-thiazol-2-yl)amino]carbonyl]amino-267430-05-7P,  
 3-[(5-Isopropyl-1,3-thiazol-2-yl)amino]carbonyl]amino-N-phenylbenzamide  
 267430-06-8P, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3-morpholinophenyl)urea  
 267430-07-9P, N-[4-[(5-Isopropyl-1,3-thiazol-2-yl)amino]carbonyl]amino-N-methylacetamide  
 267430-08-0P, N-[2-[(5-Isopropyl-1,3-thiazol-2-yl)amino]carbonyl]amino-N-[(5-  
 isopropyl-1,3-thiazol-2-yl)urea 267430-09-1P,  
 N-[2-[(5-Isopropyl-1,3-thiazol-2-yl)amino]carbonyl]amino-4-methoxyphenyl]acetamide 267430-10-4P, N-(5-Isopropyl-1,3-thiazol-2-yl)urea

2-yl)-4-(4-methoxyphenyl)-1-piperazinecarboxamide **267430-11-5P**,  
 N-(2-Furylmethyl)-N'-(5-isopropyl-1,3-thiazol-2-yl)urea  
**267430-12-6P**, N-(4-Fluorophenyl)-N'-(5-isopropyl-1,3-thiazol-2-yl)urea **267430-13-7P**, N-(2-Methoxybenzyl)-N'-(5-isopropyl-1,3-thiazol-2-yl)urea **267430-14-8P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2-(1-methyl-1H-pyrrol-2-yl)ethyl)urea **267430-15-9P**,  
 N-(3,4-Dimethoxybenzyl)-N'-(5-isopropyl-1,3-thiazol-2-yl)urea  
**267430-16-0P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-4-oxo-1-phenyl-1,3,8-triaspiro[4.5]-deca-8-carboxamide **267430-17-1P**,  
 N-(5-Isopropyl-1,3-thiazol-2-yl)-1,4-dioxa-8-azaspiro[4.5]decane-8-carboxamide **267430-18-2P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2-(1-piperidinyl)ethyl)urea **267430-19-3P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2-(4-morpholinyl)ethyl)urea **267430-20-6P**,  
 N-(4-Fluorophenyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-1-piperazinecarboxamide **267430-21-7P**, N-(4-(4-Chlorophenyl)-3-ethyl-5-isoxazol-2-yl)-N'-(5-isopropyl-1,3-thiazol-2-yl)urea  
**267430-22-8P**, 4-(4-Fluorophenyl)hydroxymethyl-N-(5-isopropyl-1,3-thiazol-2-yl)-1-piperidinecarboxamide **267430-23-9P**,  
 N-(3-Ethynylphenyl)-N'-(5-isopropyl-1,3-thiazol-2-yl)urea  
**267430-24-0P**, N-(2-Methoxy-2-fluorophenyl)-N'-(5-isopropyl-1,3-thiazol-2-yl)urea **267430-25-1P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-oxo-1-piperidinyl)urea **267430-26-2P**, N-(3-Acetylaminophenyl)-N'-(5-isopropyl-1,3-thiazol-2-yl)urea  
**267430-28-4P**, N-[4-[Ethyl(isopropyl)amino]phenyl]-N'-(5-isopropyl-1,3-thiazol-2-yl)urea **267430-29-5P**, N-(1,3-Benzodioxol-5-yl)-N'-(5-isopropyl-1,3-thiazol-2-yl)urea **267430-30-8P**,  
 N-[(5-Isopropyl-1,3-thiazol-2-yl)amino]carbonyl-amino]-1-phenyl-1H-pyrazole-4-carboxamide **267430-31-9P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-pyridinylmethyl)urea **267430-32-0P**,  
 N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-pyrazinyl)urea **267430-33-1P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(5-phenyl-1,3,4-oxadiazol-2-yl)urea  
**267430-34-2P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-4-(2-oxo-2,3-dihydro-1H-benzimidazol-1-yl)-1-piperidinecarboxamide **267430-35-3P**, N-(1,3-Benzothiazol-6-yl)-N'-(5-isopropyl-1,3-thiazol-2-yl)urea  
**267430-36-4P**, N-(1,3-Dimethyl-1H-pyrazol-5-yl)-N'-(5-isopropyl-1,3-thiazol-2-yl)urea **267430-37-5P**, N-(3-Phenyl-1-methyl-1H-pyrazol-5-yl)-N'-(5-isopropyl-1,3-thiazol-2-yl)urea **267430-38-6P**,  
 N-(5-Isopropyl-1,3-thiazol-2-yl)-3-hydroxy-1-piperidinecarboxamide  
**267430-39-7P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2-methyl-1,3-dioxo-1,3-dihydro-1H-isindol-5-yl)urea **267430-40-0P**,  
 N-(5-Isopropyl-1,3-thiazol-2-yl)-4-benzyl-1-piperazinecarboxamide  
**267430-41-1P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-4-methyl-1-piperazinecarboxamide **267430-42-2P**, 4-Hydroxy-N-(5-isopropyl-1,3-thiazol-2-yl)-1-piperidinecarboxamide **267430-43-3P**,  
 N-(5-Isopropyl-1,3-thiazol-2-yl)-3-aceticyclic[3.2.2]nonane-3-carboxamide  
**267430-44-4P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-4-(4-acetylphenyl)-1-piperazinecarboxamide **267430-45-5P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-4-oxo-2,4,5-tetrahydro-1H-1,5-benzodiazepine-1-carboxamide  
**267430-46-6P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(5,6,7,8-tetrahydro-1-naphthalenyl)urea **267430-47-7P**,  
 N-(4-Phenyl-1-thiazol-2-yl)-N'-(5-isopropyl-1,3-thiazol-2-yl)urea  
**267430-48-8P**, 4-(4-Fluorobenzyl)-N-(5-isopropyl-1,3-thiazol-2-yl)-1-piperidinecarboxamide **267430-49-9P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4,5-dihydro-2H-1,5-benzodiazepine-2-yl)urea **267430-50-2P**,  
 N-(5-Isopropyl-1,3-thiazol-2-yl)-4-(1-pyridinyl)-1-piperazinecarboxamide  
**267430-51-3P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(1H-indazol-6-yl)urea **267430-52-4P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2-chlorobenzyl)urea **267430-53-5P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(1,4-dichlorobenzyl)urea **267430-54-6P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3-fluorobenzyl)urea **267430-55-7P**,  
 N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4,4-dichlorobenzyl)urea  
**267430-56-8P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2,4-difluorobenzyl)urea **267430-57-9P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2,5-difluorobenzyl)urea **267430-58-0P**,  
 N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2,6-difluorobenzyl)urea  
**267430-59-1P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-hydroxy-3-

methoxybenzyl)urea **267430-60-4P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(5-methyl-2-furyl)urea **267430-61-5P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-methylsulfonylbenzyl)urea **267430-62-6P**, N-[(1E,2F)-2-Hydroxy-2,3-dihydro-1H-inden-1-yl]-N'-(5-isopropyl-1,3-thiazol-2-yl)urea **267430-63-7P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-chlorobenzyl)urea **267430-64-8P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2-pyridinylmethyl)urea **267430-65-9P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3,5-dimethoxybenzyl)urea **267430-66-0P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2-pyridinylmethyl)urea **267430-67-1P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-trifluoromethylbenzyl)urea **267430-68-2P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3,4,5-trimethoxybenzyl)urea **267430-69-3P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-dimethoxybenzyl)urea **267430-70-6P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-dimethylaminobenzyl)urea **267430-71-7P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2,5-dimethoxybenzyl)urea **267430-72-8P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2-chloro-6-phenoxybenzyl)urea **267430-73-9P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(1F,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)urea **267430-74-0P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3-hydroxy-4-methylphenyl)urea **267430-75-1P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-(1H-benzimidazol-2-yl)phenyl)urea **267430-77-3P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2-methyl-6-quinolinyl)urea **267430-78-4P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-(cyanomethyl)phenyl)urea **267430-79-5P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2-quinolinyl)urea **267430-80-8P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(1-oxo-2,3-dihydro-1H-inden-5-yl)urea **267430-81-9P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3-oxo-1,3-dihydro-2-benzofuran-5-yl)urea **267430-82-0P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(5-oxo-6,7,8-tetrahydro-2-naphthalenyl)urea **267430-83-1P**, Methyl 3-[[[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl]amino]-4-methylbenzoate **267430-84-2P**, Methyl 4-[[[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl]amino]-3-methylbenzoate **267430-85-3P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(4-(imidazol-1-yl)pyridin-2-yl)phenyl urea **267430-86-4P**, Ethyl 4-[[[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl]amino]benzoate **267430-87-5P**, (2R)-N-Benzyl-2-[[[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl]amino]propanamide **267430-88-6P**, 2-Hydroxy-5-[[[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl]amino]benzoic acid **267430-89-7P**, 2-Chloro-5-[[[(5-isopropyl-1,3-thiazol-2-yl)amino]carbonyl]amino]benzoic acid **267430-90-0P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(5-methyl-3-isoxazolyl)urea **267430-91-1P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(1,6-dimethoxyphenyl)urea **267430-92-2P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2,3-dimethoxybenzyl)urea **267430-93-3P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3,4-difluorobenzyl)urea **267430-94-4P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2,4-dimethylphenyl)urea **267430-95-5P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(1H-benzimidazol-2-yl)urea **267430-96-6P**, **267430-97-7P** **267430-98-8P** **267430-99-9P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(2-(1-methyl-1H-imidazol-2-yl)methoxyphenyl)urea **267431-01-6P**, N-(5-Isopropyl-1,3-thiazol-2-yl)-N'-(3-(3-methoxy-1-propynyl)phenyl)urea **267431-02-7P**, N-[3-(3-(Dimethylamino)-1-propynyl)phenyl]-N'-(5-isopropyl-1,3-thiazol-2-yl)urea **267431-03-8P**, N-[4-[[[(5-Isopropyl-1,3-thiazol-2-yl)amino]carbonyl]phenyl]methanesulfonamide **267431-04-9P**, 2-[3-[[[(5-Isopropyl-1,3-thiazol-2-yl)amino]carbonyl]amino]anilino]acetamide **267431-05-0P**, N-[3-(3-Hydroxy-1-butynyl)phenyl]-N'-(5-isopropyl-1,3-thiazol-2-yl)urea **267431-06-1P**, N-[(Imidazo[1,2-a]pyridin-2-yl)methyl]-N'-(5-isopropyl-1,3-thiazol-2-yl)urea **267431-07-2P**, N-[(5-Isopropyl-1,3-thiazol-2-yl)aminocarbonyl](2-propynyl)amino)methyl]benzenesulfonamide **267431-08-3P**, N-(1H-Indol-6-yl)-N'-(5-isopropyl-1,3-thiazol-2-yl)urea **267431-09-4P**, N-(1S)-2-Hydroxy-1-phenylethyl]-N'-(5-isopropyl-1,3-thiazol-2-yl)urea **267431-10-7P**,



DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP,  
 KE, KG, KP, KR, PS, LC, LF, LR, LS, LT, LU, LV, MD, MG, MK, MN,  
 MW, MX, NC, NZ, PL, PT, RG, EU, SD, SE, SG, SI, SK, SL, TJ, TM,  
 TF, TT, UA, UG, VE, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU,  
 TJ, TM  
 FW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,  
 ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,  
 CI, CM, GA, GN, GW, ML, MF, NE, SN, TD, TG  
 AU 9944311 A1 20000105 AU 1999-44311 19990611 --  
 EP 1087951 A1 20010404 EP 1999-927401 19990611 --  
 F: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,  
 IE, FI  
 PRAI US 1998-89747 P 19980618 --  
 WO 1999-US13034 W 19990611  
 OS MARPAT 132:35695  
 GI



AB The title compds. [I; R<sub>1</sub> = F<sub>2</sub>, COR<sub>3</sub>, CONH<sub>2</sub>, etc.; R<sub>2</sub> = alkyl, cycloalkyl, heterocycloalkyl, etc.; R<sub>3</sub> = H, alkyl, cycloalkyl, etc.; A = (CR<sub>7</sub>R<sub>8</sub>)<sub>m</sub>(CR<sub>5</sub>R<sub>6</sub>)<sub>n</sub>R<sub>4</sub> (wherein n = 0-2; m = 1-2 but both n and m cannot be 2), (CR<sub>7</sub>R<sub>8</sub>)<sub>j</sub>Y(CR<sub>5</sub>R<sub>6</sub>)<sub>i</sub>F<sub>4</sub> (i, j = 0-1 but cannot both be 1; Y = (un)substituted alkene, alkyne, any 2 adjacent carbon atoms of a cycloalkyl or cycloheteroalkyl ring of 3-7 atoms); R<sub>4</sub> = alkyl, cycloalkyl, heterocycloalkyl, etc.; F<sub>5</sub>-F<sub>8</sub> = H, alkyl, cycloalkyl, etc.], protein kinase inhibitors (no data) which are useful in the treatment of proliferative diseases, for example, cancer, inflammation, and arthritis, and also in the treatment of Alzheimer's disease, and cardiovascular disease, were prep'd. E.g., a multi-step synthesis of (E)-II, starting with 2-aminothiazol-5-ylcarboxaldehyde, was given.

IT 252660-60-9P 252660-61-0P 252660-82-5P  
 252661-05-5P 252661-06-6P 252661-07-7P  
 252661-08-8P 252661-09-9P 252661-10-2P  
 252661-11-3P 252661-12-4P 252661-13-5P  
 252661-14-6P 252661-15-7P 252661-16-8P  
 252661-17-9P 252661-18-0P 252661-19-1P  
 252661-20-4P 252661-21-5P 252661-22-6P  
 252661-23-7P 252661-24-8P 252661-25-9P  
 252661-26-0P 252661-27-1P 252661-28-2P  
 252661-29-3P 252661-31-7P 252661-32-8P  
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 252661-37-3P 252661-38-4P 252661-39-5P  
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 252661-46-4P 252661-48-6P 252661-49-7P  
 252661-54-4P 252661-55-5P 252661-56-6P  
 252661-62-4P 252661-63-5P 252661-69-1P  
 252661-70-4P 252661-71-5P 252661-97-5P  
 252662-14-9P 252662-16-1P

FL: BAC (Biological activity or effector, except adverse); SEN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PFEP Preparation); USES (Uses)  
 (prep'n. of carbon substituted aminothiazole inhibitors of cyclin dependent kinases)

IT 252660-60-9P  
 FL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PFEP

(Preparation); USES (Uses)

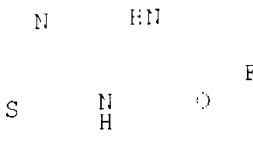
(prepn. of carbon substituted aminothiazole inhibitors of cyclin  
dependent kinases)

RN 252660-60-9 HCAPLUS

CN Urea, N-(2,6-difluorophenyl)-N'-(5-[(1E)-2-[5-(1,1-dimethylethyl)-2-  
oxazolyl]ethenyl]-2-thiazolyl)- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

F



t-Bu

RE.CNT 1

RE

(1) Boberg; US 4782162 A 1988 HCAPLUS

L26 ANSWER 4 OF 8 HCAPLUS COPYRIGHT 2001 ACS

AN 1998:776672 HCAPLUS

DN 130:38284

TI Preparation of urea derivatives as **raf** kinase inhibitorsIN Wood, Jill E.; Wild, Hanns; Rogers, Daniel H.; Lyons, John; Katz, Michael  
E.; Caringal, Yolanda V.; Dally, Robert; Lee, Wendy; Smith, Roger A.;  
Blum, Cheri L.

PA Bayer Corp., USA; Onyx Pharmaceuticals; et al.

SO PCT Int. Appl., 53 pp.

CODEM: PIXKD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 90/52554	A1	19981126	WO 1998-US10376	19980511 ---
	W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GR, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KP, KZ, LC, LF, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NE, PL, PT, EG, RU, SL, SE, SG, SI, SK, SL, TR, TM, TR, TT, UA, UG, US, US, VN, YU, ZW, AM, AE, BY, KG, KZ, MD, RU, TC, TM FW: GH, GM, KE, LS, MW, SD, SZ, UG, SW, AT, BE, CH, CY, DE, DK, ES, FI, FF, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BG, CF, CG, CI, CM, GA, GN, ML, MF, NE, SH, TD, TG				
	AU 9175855	A1	19981211	AU 1998-75855	19980511 ---
	EP 916382	A1	200006322	EP 1998-923601	19980511 ---
	E: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				

PRAI US 1997-863031 19970523 ---

WO 1998-US10376 19980521 ---

AB Substituted urea compds., useful for treating **tumors** mediated by  
**raf** kinase (no data), were prep'd. E.g., reaction of Me thioglycolate and  
3-chloro-4-methyl-2-pentenenitrile gave 16% of the 3-aminothiophene  
deriv., which was reacted with 4-MeC6H4NCO to give Me 5-isopropyl-3-(3-p-  
tolylureido)thiophene-2-carboxylate.

IT 216589-90-1P

FL: BAC (Biological activity or effector, except adverse); SPN (Synthetic  
preparation); THU (Therapeutic use); BIOL (Biological study); PREP  
(Preparation); USES (Uses)

(prepn. of urea derivs. as raf kinase inhibitors)

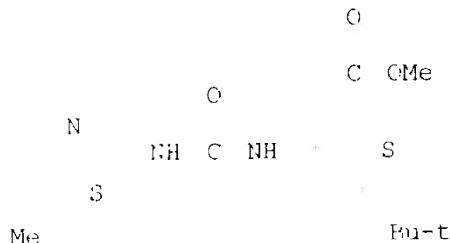
IT 216589-90-1P

FL: BAC (Biological activity or effector, except adverse); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of urea derivs. as raf kinase inhibitors)

RN 216589-90-1 HCAPLUS

CN 2-Thiophene carboxylic acid, 5-(1,1-dimethylethyl)-3-[[[(5-methyl-2-thiazolyl)amino]carbonyl]amino]-, methyl ester (9CI) (CA INDEX NAME)



RE.CNT 3

RE

- (1) Freed; US 5597719 A 1997 HCAPLUS
- (2) Kleemann; EP 676395 A2 1996 HCAPLUS
- (3) Sugen Inc; WO 96/40673 A1 1996, V87 HCAPLUS

L26 ANSWER 5 OF 8 HCAPLUS COPYRIGHT 2001 ACS

AN 1997:717901 HCAPLUS

DN 138:3680

TI Preparation of arylreas and related compounds as inhibitors of inosine 5'-monophosphate dehydrogenase.

IN Armistead, David M.; Badia, Michael C.; Bemis, Guy W.; Bethiel, Fandy S.; Frar., Catharine A.; Novak, Ferry M.; Ronkin, Steven M.; Saunders, Jeffrey D.

PA Vertex Pharmaceuticals Inc., USA

SO PCT Int. Appl., 93 pp.

CITATION: PIXXD2

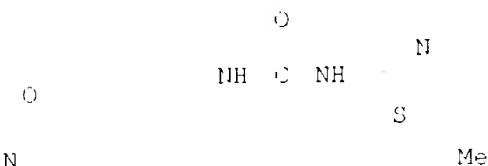
DT Patent

LA English

FAN.CNT 2

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9740038	A1	19971030	WO 1997-US6623	19970411 <--
	W: AL, AM, AT, AU, AZ, BE, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, HU, IL, IS, JP, KE, KG, KR, KZ, LC, LF, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NL, NO, PL, PT, RO, RU, SD, SE, SG, SI, SK, TJ, TM, TR, TT, UA, US, VN, YU, AM, AZ, BY, EG, IE, MD, RU, TJ, TM				
	EW: GH, KE, LS, MW, SD, SE, UG, AT, BE, CH, DE, DK, ES, FI, FR, GB, GE, IE, IT, LU, MD, NL, PT, SE, BF, BJ, CF, CG, CI, CM, CA, GN, ML, MR, NE, SN, TG, TG				
	US 5497876	A	19980915	US 1996-636361	19960423 <--
	US 6654472	A	20000415	US 1997-832165	19970402 <--
	AU 9726745	A1	19971112	AU 1997-26785	19970421 <--
	AU 7247301	B2	20000907		
	EP 9717832	A1	19990324	EP 1997-918759	19970421 <--
	F: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MS, PT, IE, SI, LT, LV, FI, KG				
	BR 9708735	A	19990813	BR 1997-8735	19970421 <--
	NC 9804917	A	19981213	NO 1998-4917	19981022 <--
PRAI	US 1996-636261	A	19960423		
	US 1997-901789	A	19971014		
	US 1997-832165	A	19970402		
	WO 1997-US6623	W	19970421		

OS MARPAT 123:3680  
 AB ANH<sub>n</sub>NHR [A = (substituted) alkyl, alkenyl, alkynyl; B = (unsatd.) (substituted) mono- or bicyclic ring contg. 1 to 4 heteroatoms; D = CO, CS, SO<sub>2</sub>], were prep'd. Thus, 4-(5-oxazolyl)aniline and PhCH<sub>2</sub>NCO were stirred overnight in CH<sub>2</sub>Cl<sub>2</sub> to give N-benzyl-N'-(4-(5-oxazolyl)phenyl)urea. Several title compds. inhibited IMPDH with Ki = 0.01-50 nM.  
 IT 198820-15-4  
 RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (prepn. of arylreas and related compds. as inhibitors of IMP dehydrogenase)  
 IT 198820-15-4  
 RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (prepn. of arylreas and related compds. as inhibitors of IMP dehydrogenase)  
 RN 198820-15-4 HCAPLUS  
 CN Urea, N-(5-methyl-2-thiazolyl)-N'-(4-(5-oxazolyl)phenyl)- (9CI) (CA INDEX NAME)



L26 ANSWER 6 OF 8 HCAPLUS COPYRIGHT 2001 ACS  
 AN 1977:50494 HCAPLUS  
 DN 86:50494  
 TI Inhibition of solid **tumors** by nitrosoureas. 1. **Lewis lung carcinoma**  
 AU Montgomery, John A.; McCaleb, George S.; Johnston, Thomas P.; Mayo, Joseph G.; Lester, W. Russell, Jr.  
 CS Kettering-Meyer Lab., South. Res. Inst., Birmingham, Ala., USA  
 SC J. Med. Chem. (1977), 20(2), 291-5  
 CODEN: JMCMAF  
 DT Journal  
 LA English  
 AB The utility of the Lewis lung **carcinoma** as a secondary screen for the evaluation of nitrosoureas as **anticancer** agents was assessed. The activity of this series of compds. was detd. against both the early (before **metastasis**) and late (after **metastasis**) forms of the disease. Although some exceptions were noted, compds. most active against the early form of the disease were most active against the established **tumor**. A differentiation in activity based on the Lewis lung system was evident with nitrosoureas equally active against leukemia L1210, although the significance of this differentiation with respect to the human disease has not yet been established.  
 IT 33024-33-8  
 RL: BAC (Biological activity or effector, except adverse); BIOL (Biological study)  
 (neoplasm inhibiting activity of)  
 IT 33024-33-8  
 RL: BAC (Biological activity or effector, except adverse); BIOL (Biological study)  
 (neoplasm inhibiting activity of)  
 RN 33024-33-8 HCAPLUS  
 CN Urea, N-(2-chloroethyl)-N-nitroso-N'-(5-nitro-2-thiazolyl)- (9CI) (CA INDEX NAME)

O NO

N NH C N CH<sub>2</sub> CH<sub>2</sub>Cl

S

O<sub>2</sub>N

L26 ANSWER 7 OF 8 HCAPLUS COPYRIGHT 2001 ACS  
 AN 1973:526436 HCAPLUS  
 DN 73:126436  
 TI Pharmaceutical 2-amino-4-aryl-5-thiazolecarboxylic acid derivatives  
 IN Manjhisi, Elso; Salimbeni, Aldo; Fregnan, Giancarlo  
 PA Istituto Luso Farmaco d'Italia S.r.l.  
 SO Ger. Offen., 20 pp.  
 CODEN: GWXXBK  
 DT Patent  
 LA German  
 FAN.CNT .

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	DE 2303251	A1	19730906	DE 1973-2109251	19730224 ---
	CA 7301203	A	19740327	ZA 1973-1308	19730220 ---
	AC 7351467	A1	19740812	AU 1973-52467	19730222 ---
	BE 795307	A1	19730618	BE 1973-118061	19730223 ---
	NL 7303528	A	19730808	NL 1973-2518	19730223 ---
	US 3333833	A	19760110	US 1973-335255	19730223 ---
	ES 412052	A1	19760501	ES 1973-412052	19730224 ---
	FR 2181764	A1	19731207	FR 1973-6736	19730226 ---
	JP 48097868	A2	19731213	JP 1973-22273	19730226 ---
	JP 51012630	B4	19760421		
	GB 1425595	A	19770218	GB 1973-9382	19730226 ---
	CA 1006515	A1	19770308	CA 1973-164590	19730226 ---

PRAI IT 1972-01086

19730225 ---

IT 1973-0231

19730209 ---

IT 1973-0231

19730209 ---

GI For diagram(s), see printed CA Issue.

AB Nine new thiazoles (I; n = 1 or 2; R = H, Cl, F, or OMe; R<sub>1</sub> = H or Et; R<sub>2</sub> = H, Et, Ac, CONHPh, Ph, or 2,6-C<sub>12</sub>C<sub>6</sub>H<sub>3</sub>; R<sub>3</sub> = OH, OEt, or NHCH<sub>2</sub>CH<sub>2</sub>NEt<sub>2</sub>) were prepd. from 4-RC<sub>6</sub>H<sub>4</sub>CO<sub>2</sub>HBr(CH<sub>2</sub>)<sub>n</sub>OEt<sub>3</sub> by reaction with H<sub>2</sub>NCSNR<sub>1</sub>R<sub>2</sub> or by reaction with H<sub>2</sub>NC(S)C<sub>2</sub>H<sub>5</sub> and subsequent chlorination and reaction with R<sub>1</sub>F<sub>2</sub>NH, from I (R<sub>3</sub> = OEt) by sapon. or reaction with Et<sub>2</sub>NCH<sub>2</sub>CH<sub>2</sub>NH<sub>2</sub>, or from I (R<sub>1</sub> = R<sub>2</sub> = H) by acetylation or reaction with PhNCO. Six I had antiinflammatory, antipyretic, antitussive, analgesic, and antitumor activity in animals and LD<sub>50</sub> 43 to >1000 and 238 to >5000 mg/kg; i.p. and orally in mice, resp.

IT 49780-01-0P 49780-06-5P

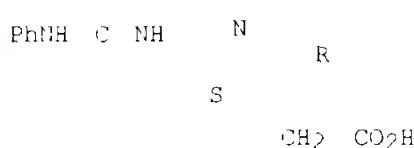
FL: SEM (Synthetic preparation); PREP (Preparation)  
(prpn. cf)

IT 49780-01-0P

FL: SEM (Synthetic preparation); PPEP (Preparation)  
(prpn. cf)

EN 49780-01-0 HCAPLUS

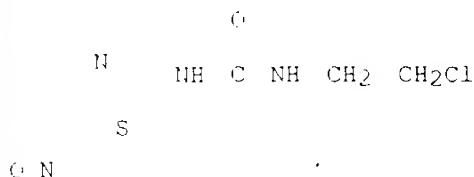
CN 5-Thiazoleacetic acid, 4-(4-chlorophenyl)-2-[(phenylamino)carbonyl]amino-  
("CI) (CA INDEX NAME)



R

Cl

L26 ANSWER 8 OF 8 HCAPLUS COPYRIGHT 2001 ACS  
 AN 1971:476218 HCAPLUS  
 DN 75:76218  
 TI Synthesis of potential **anticancer** agents. 38. N-nitrosoureas.  
 4. Further synthesis and evaluation of haloethyl derivatives  
 AU Johnston, Thomas P.; McCaler, George S.; Opliger, Pamela S.; Laster, W.  
 Russell; Montgomery, John A.  
 CS Kettering-Meyer Lab., South. Res. Inst., Birmingham, Ala., USA  
 SC J. Med. Chem. (1971), 14(7), 600-14  
 CODEN: JMCMAK  
 DT Journal  
 LA English  
 GI For diagram(s), see printed CA Issue.  
 AB N-(2-Haloethyl)-N-nitrosoureas (I), prepd. by nitrosation of the  
 corresponding 2-haloethylureas, were tested for **anticancer**  
 activity against both i.p. and intracerebrally inoculated marine leukemia  
 L1210. The chemotherapeutic indices, ED50/LD10 and ED99/LD10, were  
 compared with those of 1,3-bis(2-chloroethyl)-1-nitrosourea (II) and  
 1-(2-chloroethyl)-3-cyclohexyl-1-nitrosourea (III). 1-(2-Fluoroethyl)-1-  
 nitroso-3-(tetrahydro-2H-thiopyran-4-yl) urea S,S-dioxide,  
 1-(2-fluoroethyl)-1-nitroso-3-(tetrahydro-2H-thiopyran-4-yl)urea and  
 3-(4-acetoxy)cyclohexyl)-1-(2-chloroethyl)-1-nitrosourea were equipotent as  
 the ref. compds.  
 IT 3311-98-6P 33024-33-8P  
 RL: SPN (Synthetic preparation); PREP (Preparation)  
 (prepn. of)  
 IT 3311-98-6P  
 RL: SPN (Synthetic preparation); PREP (Preparation)  
 (prepn. of)  
 RN 3311-98-6 HCAPLUS  
 CN Urea, N-(2-chloroethyl)-N'-(5-nitro-2-thiazolyl)- (9CI) (CA INDEX NAME)



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